Detection and quantification of human colon MelQx adducts after oral MelQx administration using accelerator mass spectrometry (AMS) or ³²P-postlabelling. Mauthe R J., Dingley, K H., Leveson S H[†]., Vogel J S., Garner R C^a., Turteltaub K W. Lawrence Livermore National Laboratory, Livermore, CA 94551, ^aUniv. of York, York Y01 5DD, United Kingdom and [†]York District Hospital, York Y03 7HE, United Kingdom.

The food mutagen and carcinogen 2-amino-3,8-dimethylimidazo[4,5figuinoxaline was administered per os (MelQx, 228 µg; 370KBg/person) to two elderly human volunteers previously diagnosed with colon cancer. 4-6 Hours after ¹⁴C-MelQx ingestion, normal and tumour colon were surgically removed and a sample of blood collected. Analysis of purified colon DNA by AMS revealed a mean adduct level of ~1 adduct/ 109 bases, with tumour and normal tissue having similar adduct levels. HPLC analysis of DNA digests indicated 95% of adducts to be N-(deoxyguanosin-8-yl)-MelQx. ³²P-Postlabelling of the colon DNA samples revealed three adduct spots with similar chromatographic properties to those seen in rat liver DNA from animals dosed with MelQx. Low levels of haemoglobin adducts were detected in the blood of the dosed individuals (~20-100 pg MelQx/ g Hb). These results show that MelQx is bioavailable to human colon and that MelQx DNA adducts are formed in substantial quantities. This work was supported by the DoE (W-7405-ENG-48) من لا يسابل عليه المائلة على المائلة على المائلة المائلة على المائلة على المائلة ا and partially by NIH (CA66861), USAMRDC (MM4559FLB) and the United Kingdom MAFF (FS1722)